Amendment dated May 12, 2011 Reply to Office Action of November 12, 2010

AMENDMENTS TO THE SPECIFICATION

Please amend the following paragraph on page 895 of the Specification as follows:

Hybridization probes were generated from isolated mRNA using an Atlas TM Glass Fluorescent Labeling Kit (Clontech Laboratories, Inc., Palo Alto, California, USA), This entails a two step labeling procedure that first incorporates primary aliphatic amino groups during cDNA synthesis and then couples fluorescent dye to the cDNA by reaction with the amino NO:80420) was mixed with Poly A+ mRNA (1.5 - 2 µg mRNA isolated using the Qiagen Oligotex mRNA Spin-Column protocol or-the Stratagene Poly(A) Ouik mRNA Isolation protocol (Stratagene, La Jolla, California, USA)) in a total volume of 25 μl. The sample was incubated in a thermocycler at 70°C for 5 min, cooled to 48°C and 10 ul of 5X cDNA Synthesis Buffer (kit supplied), 5 µl 10X dNTP mix (dATP, dCTP, dGTP, dTTP and aminoallyl-dUTP; kit supplied), 7.5 µl deionized water and 2.5 µl MMLV Reverse Transcriptase (500U) added. The reaction was then incubated at 48°C for 30 minutes, followed by 1hr incubation at 42°C. At the end of the incubation the reaction was heated to 70°C for 10 min, cooled to 37°C and 0.5 µl (5 U) RNase H added, before incubating for 15 min at 37°C. The solution was vortexed for 1 min after the addition of 0.5 µl 0.5 M EDTA and 5 µl of QuickClean Resin (kit supplied) then centrifuged at 14,000-18,000 X g for 1 min. After removing the supernatant to a 0.45 µm spin filter (kit supplied), the sample was again centrifuged at 14,000-18,000 X g for 1 min, and 5.5 µl 3 M sodium acetate and 137.5 µl of 100% ethanol added to the sample before incubating at -20°C for at least 1 hr. The sample was then centrifuged at 14,000-18,000 X g at 4°C for 20 min, the resulting pellet washed with 500 µl 70% ethanol, air-dried at room temperature for 10 min and resuspended in 10 µl of 2X fluorescent labeling buffer (kit provided). 10 µl each of the fluorescent dyes Cy3 and Cy5 (Amersham Pharmacia (Piscataway, New Jersey, USA); prepared according to Atlas TM kit directions of Clontech) were added and the sample incubated in the dark at room temperature for 30 min.

Please amend the following paragraph on page 896 of the Specification as follows:

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Alternatively, 3-4 μg mRNA, 2.5 (~8.9 ng of in vitro translated mRNA) μl yeast control and 3 μg oligo dTV (TTTTTTTTTTTTTTTTTTTTTTTTT(A/C/G); -Sequence-ID-No:--X(SEQ_ID-NO:80420) were mixed in a total volume of 24.7 μl. The sample was incubated in a thermocycler at 70°C for 10 min. before chilling on ice. To this, 8 μl of 5X first strand buffer (SuperScript II RNase H- Reverse Transcriptase kit from Invitrogen (Carlsbad, California 92008); cat no. 18064022), 0.8 °C of aa-dUTP/dNTP mix (50X; 25mM dATP, 25mM dGTP, 25mM dCTP, 15mM dTTP, 10mM aminoallyl-dUTP), 4 μl of 0.1 M DTT and 2.5 μl (500 units) of Superscript R.T.II enzyme (Stratagene) were added. The sample was incubated at 42°C for 2 hours before a mixture of 10 °C of 1M NaOH and 10°C of 0.5 M EDTA were added. After a 15 minute incubation at 65°C, 25 μl of 1 M Tris pH 7.4 was added. This was mixed with 450 μl of water in a Microcon 30 column before centrifugation at 11,000 X g for 12 min. The column was washed twice with 450 μl (centrifugation at 11,000 g, 12 min.) before eluting the sample by

Please amend the following paragraph on page 900 of the Specification as follows:

inverting the Microcon column and centrifuging at 11,000 X g for 20 seconds, Sample was

Reamplification primers

99G24

CGCCAGGGTTTTCCCAGTCACGAC|ACGACTCACT|gatgagtcctgagtaal (SEO ID NO; 80421)

M13 forward

+10

MseI+0

Please amend the following paragraph on page 901 of the Specification as follows:

99G20

AGCGGATAACAATTTCACACAGGA|CACACTGGTA|tagactgcgtaccga| (SEQ ID NO:80422)

M13 reverse

+10

dehydrated by centrifugation under vacuum and stored at -20°C.

TagI +0

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Please amend the following paragraph on page 909 of the Specification as follows:

Primers used for TAIL-PCR are as follows:

AD2: 5' NGTCGASWGANAWGAA 3' (SEQ ID NO:80423) (128-fold degeneracy) S = G or C, W = A or T, and N = A, G, C, or T

Please amend the following paragraph on page 909 of the Specification as follows:

LB1: 5' GTTTAACTGCGGCTCAACTGTCT 3' (SEQ ID NO:80424)

Please amend the following paragraph on page 909 of the Specification as follows:

LB2: 5' CCCATAGACCCTTACCGCTTTAGTT 3' (SEQ ID NO:80425)

Please amend the following paragraph on page 909 of the Specification as follows:

LB3: 5' GAAAGAAAAAGAGGTATAACTGGTA 3' (SEQ ID NO:80426)

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